Amendments to the Claims

Please cancel Claims 27-30 and 40. Please amend Claims 4, 5, 41, 43 and 44. The Claim Listing below will replace all prior versions of the claims in the application:

Claim Listing

- 1. (Withdrawn) An isolated nucleic acid molecule encoding a protein with a RING-finger domain and 6 NHL-motifs wherein the protein is associated with Lafora's disease.
- 2. (Withdrawn) A nucleic acid according to Claim 1 having a sequence comprising_SEQ ID NO:1 or SEQ ID NO:3.
- 3. (Withdrawn) An isolated nucleic acid molecule according to Claim 1 comprising
 - (a) a nucleic acid sequence comprising SEQ ID NO:1 or SEQ ID NO:3, wherein T can also be U;
 - (b) a nucleic acid sequence complementary to (a);
 - (c) a nucleic acid sequence that has substantial sequence homology to a nucleic acid sequence of (a) or (b);
 - (d) a nucleic acid sequence that is an analog of a nucleic acid sequence of (a), (b) or (c); or
 - (e) a nucleic acid sequence that hybridizes to a nucleic acid sequence of (a), (b), (c) or (d) under stringent hybridization conditions.
- 4. (Currently amended) A method of detecting Lafora's disease in a mammal comprising detecting a <u>missense</u>, <u>nonsense</u> or <u>frameshift</u> mutation in a nucleic acid sequence in a sample from a mammal, wherein said nucleic acid sequence is an isolated nucleic acid molecule encoding a protein with a RING-finger domain and 6 NHL motifs, and wherein the protein is associated with Lafora's disease comprises SEQ ID NO: 1, and wherein the mutation results in a deleterious effect on the encoded protein product.

- 5. (Currently amended) A method according to Claim 4 comprising detecting a C to G change at nucleotide number 205 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 6. (Previously presented) A method according to Claim 4 comprising detecting a T to A change at nucleotide number 76 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 7. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotides GA at nucleotide positions 1048 and 1049 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 8. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotides AG at nucleotide positions 468 and 469 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 9. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotide G at nucleotide number 992 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 10. (Previously presented) A method according to Claim 4 comprising detecting a deletion of 10 bp at nucleotide positions 373 to 382 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 11. (Previously presented) A method according to Claim 4 comprising detecting a deletion of 32 bp at nucleotide positions 661 to 692 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 12. (Previously presented) A method according to Claim 4 comprising detecting a T to C change at nucleotide number 260 in the EPM2B gene sequence comprising SEQ ID NO:1.

- 13. (Previously presented) A method according to Claim 4 comprising detecting a A to C change at nucleotide number 905 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 14. (Previously presented) A method according to Claim 4 comprising detecting a T to C change at nucleotide number 98 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 15. (Previously presented) A method according to Claim 4 comprising detecting an insert of 2 Ts at nucleotide number 892 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 16. (Previously presented) A method according to Claim 4 comprising detecting a G to A change at nucleotide number 436 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 17. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotide T at nucleotide number 1100 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 18. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotide T at nucleotide position 606 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 19. (Previously presented) A method according to Claim 4 comprising detecting a A to T change at nucleotide number 923 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 20. (Previously presented) A method according to Claim 4 comprising detecting a G to T change at nucleotide number 580 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 21. (Previously presented) A method according to Claim 4 comprising detecting a G to T change at nucleotide number 199 in the EPM2B gene sequence comprising SEQ ID NO:1.

- 22. (Previously presented) A method according to Claim 4 comprising detecting a G to A change at nucleotide number 838 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 23. (Previously presented) A method according to Claim 4 comprising detecting a C to T change at nucleotide number 676 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 24. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotide A at nucleotide position 468 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 25. (Previously presented) A method according to Claim 4 comprising detecting a deletion of nucleotide C at nucleotide position 204 in the EPM2B gene sequence comprising SEQ ID NO:1.
- 26. (Previously presented) A method according to Claim 4 comprising detecting one or more mutations in the EPM2B gene as indicated in Table 1.
- 27-30. (Canceled)
- 31. (Withdrawn) A method according to Claim 4 wherein the mammal is human.
- 32. (Canceled)
- 33. (Canceled)
- 34. (Withdrawn) An isolated protein containing a RING-finger domain and six NHL domains which protein is associated with Lafora's disease.

- 35. (Withdrawn) A protein according to Claim 34 having the amino acid sequence comprising SEQ ID NO:2 or SEQ ID NO:4.
- 36. (Withdrawn) A method for detecting Lafora's disease comprising detecting a mutation in a protein according to Claim 34.
- 37. (Withdrawn) A method according to Claim 36 comprising detecting a mutation in the EPM2B protein as indicated in Table 1.
- 38. (Withdrawn) A kit for carrying out the method of Claim 4 comprising reagents for the detection of a mutation in a nucleic acid sequence comprising SEQ ID NO:1 or SEQ ID NO:3.
- 39. (Withdrawn) A kit for carrying out the method of Claim 36 comprising reagents for the detection of a mutation in a protein sequence comprising SEQ ID NO:2 or SEQ ID NO:5.
- 40. (Canceled)
- 41. (Currently amended) A method of detecting the presence or absence of Lafora's disease in a mammal human comprising detecting a mutation in the EPM2B gene nucleic acid sequence of Claim 1 wherein the nucleic acid sequence comprises:
 - (a) a nucleic acid sequence comprising SEQ ID NO:1 or SEQ ID NO:3, wherein T can also be U;
 - (b) a nucleic acid sequence complementary to (a);
 - (c) a nucleic acid sequence that has substantial sequence homology to a nucleic acid sequence of (a) or (b);
 - (d) a nucleic acid sequence that is an analog of a nucleic acid sequence of (a), (b) or (c); or
 - (e) a nucleic acid sequence that hybridizes to a nucleic acid sequence of (a), (b), (c) or (d) under stringent hybridization conditions.

- 42. (Withdrawn) A method for detecting the presence or absence of Lafora's disease comprising detecting a mutation in a protein according to claim 35.
- 43. (Currently Amended) A method of detecting the presence of absence or absence of a mutation in the nucleic acid a nucleic acid in a test sample containing the EPM2B gene sequence set forth in SEQ ID NO:1 or SEQ ID NO:3 comprising the steps of:
 - (a) analyzing a test sample containing the EPM2B gene to determine the nucleic acid sequence of the gene;
 - (b) comparing the nucleic acid sequence of the gene in the test sample to the nucleic acid sequence set forth in SEQ ID NO:1 or SEQ ID NO:3; and
 - (c) determining the differences, if any, between the sequence of the EPM2B gene in the test sample and the nucleic acid sequence set forth in SEQ ID NO:1 or SEQ ID NO:3, thereby detecting the presence of or absence of a mutation in the EPM2B gene of the test sample. nucleotide sequence set forth in SEQ ID NO:1 or SEQ ID NO:3 in a mammal.
- 44. (Currently Amended) A method for diagnosing the presence of, or predisposition to, Lafora's disease in a <u>human</u> mammal comprising:
 - (a) obtaining a nucleic acid sample from the mammal;
 - (b) analyzing <u>a</u> the nucleic acid sample <u>obtained from the human</u> to determine the presence of <u>absence of a EPM2B</u> gene mutation <u>listed in Table 1</u>, <u>associated with Lafora's disease</u>, wherein the presence of an EPM2B gene mutation <u>associated with Lafora's disease</u> indicates that the <u>human has</u>, or <u>mammal</u> is at risk for development of Lafora's disease.
- 45. (Previously presented) A method according to Claim 4 wherein the mutation is a deletion, insertion, point mutation, or repeat sequence.
- 46. (Previously presented) A method according to Claim 44 wherein the mutation is a deletion, insertion, point mutation, or repeat sequence.